SEQUENCE LISTING

(1) GENERAL	INFORMATION:
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5

(i) APPLICANT:

(A) NAME: PEPTIDE THERAPEUTICS LIMITED

(B) STREET: 100 Fulbourn Road

(C) CITY: Cambridge

10 (D) STATE: not applicable

(E) COUNTRY: United Kingdom

(F) POSTAL CODE (ZIP): CB1 9PT

(ii) TITLE OF INVENTION: ATTENUATED BACTERIA USEFUL IN VACCINES

15

(iii) NUMBER OF SEQUENCES: 6

(iv) COMPUTER READABLE FORM:

(A) MEDIUM TYPE: Floppy disk

20 (B) COMPUTER: IBM PC compatible

(C) OPERATING SYSTEM: PC-DOS/MS-DOS

(D) SOFTWARE: PatentIn Release #1.0, Version #1.30 (EPO)

(v) CURRENT APPLICATION DATA:

25 APPLICATION NUMBER:

(2) INFORMATION FOR SEQ ID NO: 1:

(i) SEQUENCE CHARACTERISTICS:

30 (A) LENGTH: 1690 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: DNA (genomic)

(vi) ORIGINAL SOURCE:

(A) ORGANISM: aroC of E.coli

(ix) FEATURE:

(A) NAME/KEY: CDS

5 (B) LOCATION: 492..1562

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

	GTCGACGCGG TGGATATCTC TCCAGACGCG CTGGCGGTTG CTGAACAGAA CATCGAAGAA	60
10	CACGGTCTGA TCCACAACGT CATTCCGATT CGTTCCGATC TGTTCCGCGA CTTGCCGAAA	120
	GTGCAGTACG ACCTGATTGT CACTAACCCG CCGTATGTCG ATGCGAAGAT ATGTCCGACC	180
	TGCCAAACAA TACCGCCACG AGCCGGAACT GGGCCTGGCA TCTGGCACTG ACGGCCTGAA	240
	ACTGACGCGT CGCATTCTCG GTAACGCGGC AGATTACCTT GCTGATGATG GCGTGTTGAT	300
	TTGTGAAGTC GGCAACAGCA TGGTACATCT TATGGAACAA TATCCGGATG TTCCGTTCAC	360
15	CTGGCTGGAG TTTGATAACG GCGGCGATGG TGTGTTTATG CTCACCAAAG AGCAGCTTAT	420
	TGCCGCACGA GAACATTTCG CGATTTATAA AGATTAAGTA AACACGCAAA CACAACAATA	480
	ACGGAGCCGT G ATG GCT GGA AAC ACA ATT GGA CAA CTC TTT CGC GTA ACC	530
	Met Ala Gly Asn Thr Ile Gly Gln Leu Phe Arg Val Thr	
	1 5 10	
20		
	ACC TTC GGC GAA TCG CAC GGG CTG GCG CTC GGC TGC ATC GTC GAT GGT	578
	Thr Phe Gly Glu Ser His Gly Leu Ala Leu Gly Cys Ile Val Asp Gly	
	15 20 25	
25	GTT CCG CCA GGC ATT CCG CTG ACG GAA GCG GAC CTG CAA CAT GAC CTC	626
	Val Pro Pro Gly Ile Pro Leu Thr Glu Ala Asp Leu Gln His Asp Leu	
	30 35 40 45	
	GAC CGT CGT CGC CCT GGG ACA TCG CGC TAT ACC ACC CAG CGC CGC GAG	674
30	Asp Arg Arg Pro Gly Thr Ser Arg Tyr Thr Thr Gln Arg Arg Glu	
	50 55 60	
	CCG GAT CAG GTC AAA ATT CTC TCC GGT GTT TTT GAA GGC GTT ACT ACC	722
	Pro Asp Gln Val Lys Ile Leu Ser Gly Val Phe Glu Gly Val Thr Thr	
35	65 70 75	

GGC ACC AGC ATT GGC TTG TTG ATC GAA AAC ACT GAC CAG CGC TCT CAG

WO 99/49026 PCT/GB99/00935

	Gly	' Thr	Ser 80	: Gly	Leu	Leu	I1e 85		Asn	Thr	Asp	G1 n 90		, Ser	Gln	
5			Ser												TAC Tyr	818
10		Tyr			TAC Tyr 115											866
i r					ACC Thr											914
15					GAG G1u											962
20					ATT Ile											1010
25					TTT Phe					Asp						1058
30					GCG Ala 195											1106
35					GCC Ala			Val					G1 y			1154
JJ	GTC Val				GAT (1202

1690



WC	77/4	7020	•														
				225					230					235			
	AAC	GCG	GTG	AAA	GGC	GTG	GAA	ATT	GGC	GAC	GGC	ПΤ	GAC	GTG	GTG	GCG	1250
	Asn	Ala	Val	Lys	Gly	Va1	Glu	Пe	G1 y	Asp	Gly	Phe	Asp	۷a٦	Val	Ala	
5			240					245					250				
_																	
	CTG	CGC	GGC	AGC	CAG	AAC	CGC	GAT	GAA	ATC	ACC	AAA	GAC	GGT	TTC	CAG	1298
	Leu	Arg	G1 y	Ser	Gln	Asn	Arg	Asp	Glu	Пе	Thr	Lys	Asp	G1 y	Phe	Gln	
		255					260					265					
10																	
	AGC	AAC	CAT	GCG	GGC	GGC	ATT	СТС	GGC	GGT	ATC	AGC	AGC	GGG	CAG	CAA	1346
	Ser	Asn	His	Ala	Gly	G1 y	Ile	Leu	Gly	Gly	Ile	Ser	Ser	Gly	Gln	Gln	
	270					275					280					285	
15	ATC	ATT	GCC	CAT	ATG	GCG	CTG	AAA	CCG	ACC	TCC	AGC	ATT	ACC	GTG	CCG	1394
	Ile	Ile	Ala	His	Met	Ala	Leu	Lys	Pro	Thr	Ser	Ser	Пe	Thr	۷a۱	Pro	
					290					295					300		
						CGC											1442
20	Gly	Arg	Thr	Ile	Asn	Arg	Phe	G1 y	Glu	Glu	Va1	Glu	Met			Lys	
				305					310					315			
																	1 400
																GAA	1490
	G1 y	Arg	His	Asp	Pro	Cys	Val			Arg	Ala	Val			Ala	Glu	
25			320					325					330				
														CCA	ACC		1538
																GGC	1556
	Ala			Gly	/ Asp	Arg			GIS	/ Ser	· Pro			Ald	1 1 1 1 1	· G1y	
		335	•				340	l				345)				
30							CAC	TCA	TAT	TCC	VCCC	TCC1	~ ^ ^ ^ ^ ^	.ΑΑ. Τ	'C A A T	ΓΔΔΔΔ(1592
									. IA	1006	いしばし	idai	~~~	vvt l	₩	ΓΑΑΑΑ	. 1002
			s cys	S Arç	y cys	G1u		,									
	350)				355	,										

35 CGCGATTGCG CTGCTGGCTC TGCTTGCCAG TAGCGCCAGC CTGGCAGCGA CGCCGTGGCA

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(2) INFORMATION FOR SEQ ID NO: 2:

(i)	SECUENCE	CHARACTERISTICS
(;)	SEUDENCE	CHARACIENTALITOS

(A) LENGTH: 356 amino acids

(B) TYPE: amino acid(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

Met Ala Gly Asn Thr Ile Gly Gln Leu Phe Arg Val Thr Thr Phe Gly
1 5 10 15

Glu Ser His Gly Leu Ala Leu Gly Cys Ile Val Asp Gly Val Pro Pro 20 25 30

Gly Ile Pro Leu Thr Glu Ala Asp Leu Gln His Asp Leu Asp Arg Arg
35 40 45

Arg Pro Gly Thr Ser Arg Tyr Thr Thr Gln Arg Arg Glu Pro Asp Gln
50 55 60

Val Lys Ile Leu Ser Gly Val Phe Glu Gly Val Thr Thr Gly Thr Ser
25 65 70 75 80

Ile Gly Leu Leu Ile Glu Asn Thr Asp Gln Arg Ser Gln Asp Tyr Ser 85 90 95

30 Ala Ile Lys Asp Val Phe Arg Pro Gly His Ala Asp Tyr Thr Tyr Glu 100 105 110

Gln Lys Tyr Gly Leu Arg Asp Tyr Arg Gly Gly Gly Arg Ser Ser Ala 115 120 125

Arg Glu Thr Ala Met Arg Val Ala Ala Gly Ala Ile Ala Lys Lys Tyr 130 135 140

	Leu A	la '	Glu	Lys	Phe	Gly 150	lle	G1u	Ile	Arg	G1y 155	Cys	Leu	Thr		Met 160
5	Gly A	lsp	Ile	Pro	Leu 165	Asp	Пе	Lys	Asp	Trp 170	Ser	Gln	Val		G1n 175	Asn
	Pro f	he	Phe	Cys 180	Pro	Asp	Pro	Asp	Lys 185	Ile	Asp	Ala	Leu	Asp 190	Glu	Leu
10	Met /	Arg	Ala 195	Leu	Lys	Lys	Glu	Gly 200	Asp	Ser	Ile	Gly	A1 a 205	Lys	Val	Thr
	Val '	Val 210	Ala	Ser	Gly	Val	Pro 215	Ala	Gly	Leu	Gly	G1u 220	Pro	Val	Phe	Asp
15	Arg 225	Leu	Asp	Ala	Asp	Ile 230	Ala	His	Ala	Leu	Met 235		Ile	Asn	Ala	Va1 240
20	Lys	G1 y	Va1	Glu	Ile 245	Gly	Asp	Gly	Phe	250		Val	Ala	Leu	Arg 255	
	Ser	Gln	Asr	a Arg 260		G Tu	Ile	. Thr	Lys 265		o Gly	/ Phe	e Glr	Ser 270		His
25	Ala	G٦χ	/ G1 y 275		e Lei	ı Gly	Gly	/ Ile 280		- Sei	r Gly	y G1:	n G1r 289		e Ile	e Ala
	His	Me1		a Lei	u Ly:	s Pro	7hi 29!		r Sei	r Il	e Th	r Va 30		o Gly	y Arg	g Thr
30	I1e 305		n Ar	g Ph	e G1	y G1:		u Va	1 G1	u Me	t Il		r Ly	s G1;	ý Ar	g His 320

35 Asp Pro Cys Val Gly Ile Arg Ala Val Pro Ile Ala Glu Ala Asn Ala

325

330



G1y	Asp	Arg	Phe	Asn	Gly	Ser	Pro	Val	Thr	Ala	Thr	Gly	Ala	Lys	Cys
			340					345					350		

Arg	Cys	Glu	Asp	*
		355		

(2) INFORMATION FOR SEQ ID NO: 3:

		OULD A OTED TOTTOC
(1)	SEQUENCE	CHARACTERISTICS:

10 (A) LENGTH: 1713 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: DNA (genomic)

(vi) ORIGINAL SOURCE:

(A) ORGANISM: ompC of E.coli

20 (ix) FEATURE:

30

35

(A) NAME/KEY: CDS

(B) LOCATION: 491..1594

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

GTTAACAAGC GTTATAGTTT TTCTGTGGTA GCACAGAATA ATGAAAAGTG TGTAAAGAAG 60

GGTAAAAAAA ACCGAATGCG AGGCATCCGG TTGAAATAGG GGTAAACAGA CATTCAGAAA 120

TGAATGACGG TAATAAATAA AGTTAATGAT GATAGCGGGA GTTATTCTAG TTGCGAGTGA 180

AGGTTTTGTT TTGACATTCA GTGCTGTCAA ATACTTAAGA ATAAGTTATT GATTTTAACC 240

TTGAATTATT ATTGCTTGAT GTTAGGTGCT TATTTCGCCA TTCCGCAATA ATCTTAAAAA 300

GTTCCCTTGC ATTTACATTT TGAAACATCT ATAGCGATAA ATGAAAACATC TTAAAAGTTT 360

PCT/GB99/00935 WO 99/49026 TAGTATCATA TTCGTGTTGG ATTATTCTGC ATTTTTGGGG AGAATGGACT TGCCGACTGA TTAATGAGGG TTAATCAGTA TGCAGTGGCA TAAAAAAGCA AATAAAGGCA TATAACAGAG GGTTAATAAC ATG AAA GTT AAA GTA CTG TCC CTG GTC CCA GCT CTG Met Lys Val Lys Val Leu Ser Leu Leu Val Pro Ala Leu CTG GTA GCA GGC GCA GCA AAC GCT GCT GAA GTT TAC AAC AAA GAC GGC Leu Val Ala Gly Ala Ala Asn Ala Glu Val Tyr Asn Lys Asp Gly AAC AAA TTA GAT CTG TAC GGT AAA GTA GAC GGC CTG CAC TAT TTC TCT Asn Lys Leu Asp Leu Tyr Gly Lys Val Asp Gly Leu His Tyr Phe Ser GAC AAC AAA GAT GTA GAT GGC GAC CAG ACC TAC ATG CGT CTT GGC TTC Asp Asn Lys Asp Val Asp Gly Asp Gln Thr Tyr Met Arg Leu Gly Phe AAA GGT GAA ACT CAG GTT ACT GAC CAG CTG ACC GGT TAC GGC CAG TGG Lys Gly Glu Thr Gln Val Thr Asp Gln Leu Thr Gly Tyr Gly Gln Trp GAA TAT CAG ATC CAG GGC AAC AGC GCT GAA AAC GAA AAC AAC TCC TGG Glu Tyr Gln Ile Gln Gly Asn Ser Ala Glu Asn Glu Asn Asn Ser Trp ACC CGT GTG GCA TTC GCA GGT CTG AAA TTC CAG GAT GTG GGT TCT TTC Thr Arg Val Ala Phe Ala Gly Leu Lys Phe Gln Asp Val Gly Ser Phe GAC TAC GGT CGT AAC TAC GGC GTT GTT TAT GAC GTA ACT TCC TGG ACC 35 Asp Tyr Gly Arg Asn Tyr Gly Val Val Tyr Asp Val Thr Ser Trp Thr

PCT/GB99/00935 WO 99/49026 GAC GTA CTG CCA GAA TTC GGT GGT GAC ACC TAC GGT TCT GAC AAC TTC Asp Val Leu Pro Glu Phe Gly Gly Asp Thr Tyr Gly Ser Asp Asn Phe ATG CAG CAG CGT GGT AAC GGC TTC GCG ACC TAC CGT AAC ACT GAC TTC ·Met Gln Gln Arg Gly Asn Gly Phe Ala Thr Tyr Arg Asn Thr Asp Phe TTC GGT CTG GTT GAC GGC CTG AAC TTT GCT GTT CAG TAC CAG GGT AAA Phe Gly Leu Val Asp Gly Leu Asn Phe Ala Val Gln Tyr Gln Gly Lys AAC GGC AAC CCA TCT GGT GAA GGC TTT ACT AGT GGC GTA ACT AAC AAC Asn Gly Asn Pro Ser Gly Glu Gly Phe Thr Ser Gly Val Thr Asn Asn GGT CGT GAC GCA CTG CGT CAA AAC GGC GAC GGC GTC GGC GGT TCT ATC Gly Arg Asp Ala Leu Arg Gln Asn Gly Asp Gly Val Gly Gly Ser Ile ACT TAT GAT TAC GAA GGT TTC GGT ATC GGT GGT GCG ATC TCC AGC TCC Thr Tyr Asp Tyr Glu Gly Phe Gly Ile Gly Gly Ala Ile Ser Ser Ser AAA CGT ACT GAT GCT CAG AAC ACC GCT GCT TAC ATC GGT AAC GGC GAC Lys Arg Thr Asp Ala Gln Asn Thr Ala Ala Tyr Ile Gly Asn Gly Asp CGT GCT GAA ACC TAC ACT GGT GGT CTG AAA TAC GAC GCT AAC AAC ATC Arg Ala Glu Thr Tyr Thr Gly Gly Leu Lys Tyr Asp Ala Asn Asn Ile - 610 TAC CTG GCT GCT CAG TAC ACC CAG ACC TAC AAC GCA ACT CGC GTA GGT Tyr Leu Ala Ala Gln Tyr Thr Gln Thr Tyr Asn Ala Thr Arg Val Gly TCC CTG GGT TGG GCG AAC AAA GCA CAG AAC TTC GAA GCT GTT GCT CAG

	Ser	Leu	Gly	Trp 630	Ala	Asn	Lys	Ala	G1n 635	Asn	Phe	Glu	Ala	Va1 640	Ala	Gln	
	TAC	CAG	ттс	GAC	ттс	GGT	CTG	CGT	CCG	TCC	CTG	GCT	TAC	CTG	CAG	TCT	1393
5	Tyr	Gln	Phe	Asp	Phe	Gly	Leu	Arg	Pro	Ser	Leu	Ala	Tyr	Leu	Gln	Ser	
			645					650					655				
															070		1.441
			AAA														1441
	Lys		Lys	Asn	Leu	Gly		Gly	Tyr	Asp	Asp		ASP	rie	Leu	Lys	
10		660					665					670					
	TAT	GTT	GAT	GTT	GGT	GCT	ACC	TAC	TAC	ттс	AAC	AAA	AAC	ATG	TCC	ACC	1489
			Asp														
	675					680					685					690	
15																	
	TAC	GTT	GAC	TAC	AAA	ATC	AAC	CTG	CTG	GAC	GAC	AAC	CAG	TTC	ACT	CGT	1537
	Tyr	۷a٦	Asp	Tyr	Lys	Ile	Asn	Leu	Leu	Asp	Asp	Asn	Gln	Phe		Arg	
					695					700					705		
									ATC	CTA	ССТ	· cTC	CCT	CTC	стт	TAC	1585
20																TAC Tyr	2000
	Asp	АІа	ыц	710		1 1111	Mah	ASII	715		Αι σ			720			
				710	,				0					•			
	CAG	TTC	TAA	тст	CGAT	TGA	TATO	GAAC	AA G	GGCC	TGCG	G GC	сстт	1111	-		1634
25	G1n	Phe	*														
			725	,													
	CAT	TGTT	ттс	AGC	STAC/	AAA (CTCAG	m	T T(GGTG	ract(сто	GCGA	CCGT	TCGC	CATGAGG	1694
30	ATA	VATC/	ACGT	ACG	AAA?	TΑ											1713
50	,	;									=	··					

(2) INFORMATION FOR SEQ ID NO: 4:

(i) SEQUENCE CHARACTERISTICS:

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(A) LENGTH: 367 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii)	MOLECULE	TYPE:	protein
(' ' ' '	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,		•

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

5

Met Lys Val Lys Val Leu Ser Leu Leu Val Pro Ala Leu Leu Val Ala 1 5 10 15

Gly Ala Ala Asn Ala Ala Glu Val Tyr Asn Lys Asp Gly Asn Lys Leu 10 20 25 30

Asp Leu Tyr Gly Lys Val Asp Gly Leu His Tyr Phe Ser Asp Asn Lys
35 40 45

15 Asp Val Asp Gly Asp Gln Thr Tyr Met Arg Leu Gly Phe Lys Gly Glu
50 55 60

Thr Gln Val Thr Asp Gln Leu Thr Gly Tyr Gly Gln Trp Glu Tyr Gln
65 70 75 80

20

Ile Gln Gly Asn Ser Ala Glu Asn Glu Asn Asn Ser Trp Thr Arg Val 85 90 95

Ala Phe Ala Gly Leu Lys Phe Gln Asp Val Gly Ser Phe Asp Tyr Gly
100 105 110

Arg Asn Tyr Gly Val Val Tyr Asp Val Thr Ser Trp Thr Asp Val Leu
115 120 125

Pro Glu Phe Gly Gly Asp Thr Tyr Gly Ser Asp Asn Phe Met Gln Gln
130 135 140

Arg Gly Asn Gly Phe Ala Thr Tyr Arg Asn Thr Asp Phe Phe Gly Leu 145 150 155 160

35

Val Asp Gly Leu Asn Phe Ala Val Gln Tyr Gln Gly Lys Asn Gly Asn 165 170 175

35

355

	Pro	Ser	G1y	Glu 180		Phe	Thr	Ser	Gly 185		Thr	Asn	Asn	Gly 190	-	Asp
5	Ala	Leu	Arg 195		Asn	G1 y	Asp	G1 <i>y</i> 200		Gly	Gly	Ser	Ile 205	Thr	Tyr	Asp
	Tyr	G1u 210	Gly	Phe	Gly	Ile	Gly 215	Gly	Ala	Ile	Ser	Ser 220	Ser	Lys	Arg	Thr
10	Asp 225	Ala	Gln	Asn	Thr	A1a 230	Ala	Tyr	Ile	Gly	Asn 235	Gly	Asp	Arg	Ala	G1u 240
15	Thr	Tyr	Thr	Gly	G1 y 245	Leu	Lys	Tyr	Asp	A1a 250	Asn	Asn	Ile	Tyr	Leu 255	Ala
	Ala	Gln	Tyr	Thr 260	Gln	Thr	Tyr	Asn	A1 a 265	Thr	Arg	Val	Gly	Ser 270	Leu	Gly
20	Trp	Ala	Asn 275	Lys	Ala	Gln	Asn	Phe 280	G1u	Ala	Val	Ala	G1n 285	Tyr	Gln	Phe
	Asp	Phe 290	Gly	Leu	Arg		Ser 295	Leu	Ala	Tyr	Leu	G1n 300	Ser	Lys	Gly	Lys
25	Asn 305	Leu	G1 y	Arg	Gly	Tyr 310	Asp	Asp	Glu	Asp	Ile 315	Leu	Lys	Tyr		Asp 320
30	Val	Gly	Ala	Thr	Tyr 325	Tyr	Phe	Asn		Asn 330	Met	Ser	Thr		Va1 335	Asp
	Tyr	Lys		Asn 340	Leu	Leu .	Asp	•	Asn 345	G1n	Phe	Thr .	. •	Asp 350	Ala (G1 <i>y</i>

365

Ile Asn Thr Asp Asn Ile Val Ala Leu Gly Leu Val Tyr Gln Phe *



(2) INFORMATION FOR SEQ ID NO	: 5	5	5
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(i) SEQUENCE CHARACTERIST	Ίi	i) SEC)UENCE	CHARACT	ERISTI	CS:
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(A) LENGTH: 1808 base pairs

5 (B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

10

(vi) ORIGINAL SOURCE:

(A) ORGANISM: ompF of E.coli

(ix) FEATURE:

15 (A) NAME/KEY: CDS

(B) LOCATION: 457..1545

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

20

35

AAAACTAATC CGCATTCTTA TTGCGGATTA GTTTTTTCTT AGCTAATAGC ACAATTTTCA 60 TACTATTTT TGGCATTCTG GATGTCTGAA AGAAGATTTT GTGCCAGGTC GATAAAGTTT 120 25 CCATCAGAAA CAAAATTTCC GTTTAGTTAA TTTAAATATA AGGAAATCAT ATAAATAGAT 180 TAAAATTGCT GTAAATATCA TCACGTCTCT ATGGAAATAT GACGGTGTTC ACAAAGTTCC 240 TTAAATTITA CTTTTGGTTA CATATTTTTT CTTTTTGAAA CCAAATCTTT ATCTTTGTAG 300 30 CACTITCACG GTAGCGAAAC GTTAGTTTGA ATGGAAAGAT GCCTGCAGAC ACATAAAGAC 360 ACCAAACTCT CATCAATAGT TCCGTAAATT TTTATTGACA GAACTTATTG ACGGCAGTGG 420

CAGGTGTCAT AAAAAAACC ATGAGGGTAA TAAATA ATG ATG AAG CGC AAT ATT

Met Met Lys Arg Asn Ile

1

WC	99/49026	
	CTG GCA GTG ATC GTC CCT GCT CTG TTA GTA GCA GGT ACT GCA AAC GCT Leu Ala Val Ile Val Pro Ala Leu Leu Val Ala Gly Thr Ala Asn Ala 10 15 20	522
5	GCA GAA ATC TAT AAC AAA GAT GGC AAC AAA GTA GAT CTG TAC GGT AAA Ala Glu Ile Tyr Asn Lys Asp Gly Asn Lys Val Asp Leu Tyr Gly Lys 25 30 35	570
10	GCT GTT GGT CTG CAT TAT TTT TCC AAG GGT AAC GGT GAA AAC AGT TAC Ala Val Gly Leu His Tyr Phe Ser Lys Gly Asn Gly Glu Asn Ser Tyr 40 45 50	618
15	GGT GGC AAT GGC GAC ATG ACC TAT GCC CGT CTT GGT TTT AAA GGG GAA Gly Gly Asn Gly Asp Met Thr Tyr Ala Arg Leu Gly Phe Lys Gly Glu 55 60 65 70	666
	ACT CAA ATC AAT TCC GAT CTG ACC GGT TAT GGT CAG TGG GAA TAT AAC Thr Gln Ile Asn Ser Asp Leu Thr Gly Tyr Gly Gln Trp Glu-Tyr Asn 75 80 85	714
20	TTC CAG GGT AAC AAC TCT GAA GGC GCT GAC GCT CAA ACT GGT AAC AAA Phe Gin Gly Asn Asn Ser Glu Gly Ala Asp Ala Gin Thr Gly Asn Lys 90 95 100	762
25	ACG CGT CTG GCA TTC GCG GGT CTT AAA TAC GCT GAC GTT GGT TCT TTC Thr Arg Leu Ala Phe Ala Gly Leu Lys Tyr Ala Asp Val Gly Ser Phe 105 110 115	810
30	GAT TAC GGC CGT AAC TAC GGT GTG GTT TAT GAT GCA CTG GGT TAC ACC Asp Tyr Gly Arg Asn Tyr Gly Val Val Tyr Asp Ala Leu Gly Tyr Thr 120 125 130	858
35	GAT ATG CTG CCA GAA TTT GGT GGT GAT ACT GCA TAC AGC GAT GAC TTC Asp Met Leu Pro Glu Phe Gly Gly Asp Thr Ala Tyr Ser Asp Asp Phe 135 140 145 150	906
	TTC GTT GGT CGT GTT GGC GGC GTT GCT ACC TAT CGT AAC TCC AAC TTC	954



	Phe	Val	Gly		Val 155	Gly	Gly	Val	Ala	Thr 160	Tyr	Arg	Asn	Ser	Asn 165	Phe	
5															GGT Gly		1002
				170					175					180			
	AAC	GAG	CGT	GAC	ACT	GCA	CGC	CGT	тст	AAC	GGC	GAC	GGT	GTT	GGC	GGT	1050
															Gly		
10	7311	4.0	185				J	190					195				
10																	
	тст	ATC	AGC	TAC	GAA	TAC	GAA	GGC	Ш	GGT	ATC	GTT	GGT	GCT	TAT	GGT	1098
	Ser	Ιle	Ser	Tyr	Glu	Tyr	Glu	G1 y	Phe	G1 y	Ile	Val	Gly	Ala	Tyr	Gly	
		200					205	-				210					
15																	
															AAC		1146
	Ala	Ala	Asp	Arg	Thr	Asn	Leu	G1n	Glu	Ala	Gln	Pro	Leu	Gly	Asn		
	215					220					225					230	
								_								440	1104
20															AAC		1194
	Lys	Lys	Ala	Glu			Ala	Thr	Gly		Lys	ıyr	ASP	Ala	Asn	ASII	
					235					240					245		
	ATC	TAC	CTC	CCA	ccc	۸۸۲	TAC	GGT	GAA	ACC	CGT	AAC	GCT	ACG	CCG	ATC	1242
25															Pro		
25	116	1 91	Leu	250		7311		۵.,	255	••••	3			260			
				230													
	ACT	AAT	AAA	П	ACA	AAC	ACC	AGC	GGC	ттс	GCC	AAC	AAA	ACG	CAA	GAC	1290
																Asp	
30			265	ı				270					275				
											\$						

GTT CTG TTA GTT GCG CAA TAC CAG TTC GAT TTC GGT CTG CGT CCG TCC

1338

Val Leu Leu Val Ala Gln Tyr Gln Phe Asp Phe Gly Leu Arg Pro Ser

280

285

290

ATC GCT TAC ACC AAA TCT AAA GCG AAA GAC GTA GAA GGT ATC GGT GAT

1386

Ile Ala Tyr Thr Lys Ser Lys Ala Lys Asp Val Glu Gly Ile Gly Asp



wo	99/4	9026														•	PCT/GB99/
	295					300					305					310	
			CTG														1434
5	Val	Asp	Leu	Val	Asn 315	Tyr	Phe	Glu	Val	G1 y 320	Ala	Thr	Tyr	Tyr	Phe 325	Asn	
	AAA	AAC	ATG	TCC	ACC	TAT	GTT	GAC	TAC	ATC	ATC	AAC	CAG	ATC	GAT	TCT	1482
	Lys	Asn	Met	Ser 330	Thr	Tyr	Val	Asp	Tyr 335	Ile	Ile	Asn	Gln	11e 340	Asp	Ser	
10											400		007	CTC	ОСТ	ATC	1530
			AAA Lys														1530
	ASP	ASII	345	Leu	diy	Vui	uij	350	ЛЭР	ЛОР			355				
15	-		CAG Gln			TAG	CACA	CCT (СТТТ	GTTA	AA T	GCCG	AAA A	A AC	AGGA	стт	1585
	GGT	CCTG		Ш	TATA	сс т	TCCA	GAGC	A AT	CTCA	CGTC	TTG	CAAA	AAC .	AGCC	TGCGT	T 1645
20	ттс	ATCA	GTA	ATAG	TTGG	AA T	тπс	TAAA	т ст	CCCG	TTAC	ССТ	GATA	GCG	GACT	тссст	T 1705
	CTG	TAAC	CAT	AATG	GAAC	ст с	GTCA	тстт	T GA	GAAC	ATTA	CCG	CCGC	тсс	TGCC	GACCC	G 1765
25	ATT	CTGG	GCC	TGGC	CGAT	CT G	ттс	GTGC	C GA	TGAA	CGTC	CCG	i				1808
	(2)	INF	ORMA	NOIT	l FOR	SEC	ID	NO:	6:								
30			(i)	SEQL	JENCE	CHA	RACT	ERIS	TICS	: :							
		5								4					3		

- (A) LENGTH: 362 amino acids.
- (B) TYPE: amino acid
- (D) TOPOLOGY: linear
- 35 (ii) MOLECULE TYPE: protein
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:



Met	Met	Lys	Arg	Asn	Пe	Leu	Ala	۷a۱	Пe	Val	Pro	Ala	Leu	Leu	Val
1				5					10					15	

- Ala Gly Thr Ala Asn Ala Ala Glu Ile Tyr Asn Lys Asp Gly Asn Lys
 5 20 25 30
 - Val Asp Leu Tyr Gly Lys Ala Val Gly Leu His Tyr Phe Ser Lys Gly
 35 40 45
- Asn Gly Glu Asn Ser Tyr Gly Gly Asn Gly Asp Met Thr Tyr Ala Arg
 50 55 60
 - Leu Gly Phe Lys Gly Glu Thr Gln Ile Asn Ser Asp Leu Thr Gly Tyr
 65 70 75 80
- Gly Gln Trp Glu Tyr Asn Phe Gln Gly Asn Asn Ser Glu Gly Ala Asp
 85 90 95
- Ala Gln Thr Gly Asn Lys Thr Arg Leu Ala Phe Ala Gly Leu Lys Tyr
 20 100 105 110
 - Ala Asp Val Gly Ser Phe Asp Tyr Gly Arg Asn Tyr Gly Val Val Tyr
 115 120 125
- 25 Asp Ala Leu Gly Tyr Thr Asp Met Leu Pro Glu Phe Gly Gly Asp Thr 130 135 140
 - Ala Tyr Ser Asp Asp Phe Phe Val Gly Arg Val Gly Gly Val Ala Thr 145 150 155 160
- Tyr Arg Asn Ser Asn Phe Phe Gly Leu Val Asp Gly Leu Asn Phe Ala
 165 170 175
- Val Gln Tyr Leu Gly Lys Asn Glu Arg Asp Thr Ala Arg Arg Ser Asn 35 180 185 190
 - Gly Asp Gly Val Gly Gly Ser Ile Ser Tyr Glu Tyr Glu Gly Phe Gly



195 200 205

Ile Val Gly Ala Tyr Gly Ala Ala Asp Arg Thr Asn Leu Gln Glu Ala 210 215 220

- Gln Pro Leu Gly Asn Gly Lys Lys Ala Glu Gln Trp Ala Thr Gly Leu 225 230 235 240
- Lys Tyr Asp Ala Asn Asn Ile Tyr Leu Ala Ala Asn Tyr Gly Glu Thr

 245 250 255
 - Arg Asn Ala Thr Pro Ile Thr Asn Lys Phe Thr Asn Thr Ser Gly Phe
 260 265 270
- 15 Ala Asn Lys Thr Gln Asp Val Leu Leu Val Ala Gln Tyr Gln Phe Asp 275 280 285
 - Phe Gly Leu Arg Pro Ser Ile Ala Tyr Thr Lys Ser Lys Ala Lys Asp 290 295 300
- Val Glu Gly Ile Gly Asp Val Asp Leu Val Asn Tyr Phe Glu Val Gly 305 310 315 320
- Ala Thr Tyr Tyr Phe Asn Lys Asn Met Ser Thr Tyr Val Asp Tyr Ile
 25 325 330 335
 - Ile Asn Gln Ile Asp Ser Asp Asn Lys Leu Gly Val Gly Ser Asp Asp 340 345 350
- 30 Thr Val Ala Val Gly Ile Val Tyr Gln Phe * 355 360